FULL ESTIMATED COST

7.48 51.06

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STRUCTURE FILE UPDATES: 29 MAY 2002 HIGHEST RN 423115-51-9 DICTIONARY FILE UPDATES: 29 MAY 2002 HIGHEST RN 423115-51-9

TSCA INFORMATION NOW CURRENT THROUGH July 7, 2001

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> s flunarizine or 122 L23 2 FLUNARIZINE OR L22

=> s calcium L24 70643 CALCIUM

=> fil caplus
COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL ENTRY SESSION 8.38 59.44

FILE 'CAPLUS' ENTERED AT 22:09:46 ON 30 MAY 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 30 May 2002 VOL 136 ISS 22 FILE LAST UPDATED: 29 May 2002 (20020529/ED)

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=> s calcium channel and 126 L27 372 CALCIUM CHANNEL AND L26

=> s (calcium channel blocker and 126 and skin UNMATCHED LEFT PARENTHESIS '(CALCIUM' The number of right parentheses in a query must be equal to the number of left parentheses.

 , thesaurus (/RL field) in this file.

=> s flunarizine or 122

1168 FLUNARIZINE OR L22 L26

=> s calcium channel and 126

372 CALCIUM CHANNEL AND L26

=> s (calcium channel blocker and 126 and skin UNMATCHED LEFT PARENTHESIS '(CALCIUM' The number of right parentheses in a query must be equal to the number of left parentheses.

=> s (calcium channel blocker and 126 and skin) 2 (CALCIUM CHANNEL BLOCKER AND L26 AND SKIN)

=> d 1-2 all

=> Uploading 09981751.str

L2 STRUCTURE UPLOADED

=> d 12

L2 HAS NO ANSWERS

Structure attributes must be viewed using STN Express query preparation.

=> s 12 sss ful

FULL SEARCH INITIATED 22:44:59

FULL SCREEN SEARCH COMPLETED -202 TO ITERATE

100.0% PROCESSED 202 ITERATIONS

SEARCH TIME: 00.00.01

L3 68 SEA SSS FUL L2

=> fil caplus

SINCE FILE COST IN U.S. DOLLARS TOTAL ENTRY SESSION 157.04 FULL ESTIMATED COST 156.83

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL

ENTRY SESSION

68 ANSWERS

-CA GUBSCRIBER PRICE

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FILE COVERS 1907 - 30 May 2002 VOL 136 ISS 22 FILE LAST UPDATED: 29 May 2002 (20020529/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> s 13 and calcium channel blocker L4 179 L3 AND CALCIUM CHANNEL BLOCKER

=> s 14 and skin

L5 2 L4 AND SKIN

=> d 1-2 all

(FILE 'HOME' ENTERED AT 22:37:59 ON 30 MAY 2002)

FILE 'REGISTRY' ENTERED AT 22:38:07 ON 30 MAY 2002

L1 1 S FLUNARIZINE HYDROCHLORIDE

L2 STRUCTURE UPLOADED

L3 68 S L2 SSS FUL

FILE 'CAPLUS' ENTERED AT 22:45:05 ON 30 MAY 2002

179 S L3 AND CALCIUM CHANNEL BLOCKER

L5 2 S L4 AND SKIN

=> log y

L4

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 22.19 179.23

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE -2.48 -3.07

STN INTERNATIONAL LOGOFF AT 22:46:40 ON 30 MAY 2002

29 MAY 2002 HIGHEST RN 423115-51-9 STRUCTURE FILE UPDATES: DICTIONARY FILE UPDATES: 29 MAY 2002 HIGHEST RN 423115-51-9 TSCA INFORMATION NOW CURRENT THROUGH July 7, 2001 Please note that search-term pricing does apply when conducting SmartSELECT searches. Crossover limits have been increased. See HELP CROSSOVER for details. Calculated physical property data is now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf => s flunarizine hydrochloride 1 FLUNARIZINE HYDROCHLORIDE => d all ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS T.1 30484-77-6 REGISTRY
Piperazine, 1-[bis(4-fluorophenyl)methyl]-4-[(2E)-3-phenyl-2-propenyl]-, RN dihydrochloride (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: Piperazine, 1-[bis(4-fluorophenyl)methyl]-4-(3-phenyl-2-propenyl)-, dihydrochloride, (E)-Piperazine, 1-[bis(p-fluorophenyl)methyl]-4-cinnamyl-, dihydrochloride, CN (E) - (8CI)OTHER NAMES: Flunarizine dihydrochloride CN Flunarizine hydrochloride CN R 14950 CN FS STEREOSEARCH C26 H26 F2 N2 . 2 Cl H MF CT ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO, CA, CAPLUS, CHEMCATS, LC STN Files: CHEMLIST, CIN, CSCHEM, DRUGPAT, DRUGUPDATES, EMBASE, IPA, MRCK*, PHAR, RTECS*, TOXCENTER, USAN, USPATFULL (*File contains numerically searchable property data) EINECS** Other Sources: (**Enter CHEMLIST File for up-to-date regulatory information) CRN (52468-60-7)Ring System Data Elemental|Elemental| Size of |Ring System| Ring | RID Analysis |Sequence | the Rings | Formula | Identifier | Occurrence | ES | SZ | RF | RID | Count _____+__+__+__+ C6 1C6 16 [C6 |46.150.18 |3 C4N2 - NC2NC2 16 |C4N2 |46.383.1 |1

Double bond geometry as shown.

●2 HC1

38 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
38 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1

Uploading 09981751.str

L2 STRUCTURE UPLOADED

=> d 12

L2 HAS NO ANSWERS

L2 STR

Structure attributes must be viewed using STN Express query preparation.

=> s 12 sss ful

FULL SEARCH INITIATED 22:44:59

FULL SCREEN SEARCH COMPLETED - 2

202 TO ITERATE

100.0% PROCESSED

202 ITERATIONS

68 ANSWERS

SEARCH TIME: 00.00.01

L3 68 SEA SSS FUL L2

=> fil caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION

FULL ESTIMATED COST 156.83 157.04

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE -0.59 -0.59

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FILE COVERS 1907 - 30 May 2002 VOL 136 ISS 22 FILE LAST UPDATED: 29 May 2002 (20020529/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file. => s 13 and calcium channel blocker 179 L3 AND CALCIUM CHANNEL BLOCKER => s 14 and skin 2 L4 AND SKIN => d 1-2 all ANSWER 1 OF 2 CAPLUS COPYRIGHT 2002 ACS 1999:339238 CAPLUS ΑN 131:139372 DN Effects of the calcium channel blocker TIflunarizine on spinal reflexes in cats Genc, O.; Demir, S.; Tasci, N.; Kaptanoglu, B.; Marangoz, C. ΑIJ Department of Physiology, Faculty of Medicine, Pamukkale University, CS Denizli, 20070, Turk. Medical Science Research (1999), 27(4), 247-250 SO CODEN: MSCREJ; ISSN: 0269-8951 PΒ Lippincott Williams & Wilkins DTJournal English LA CC 1-11 (Pharmacology) Section cross-reference(s): 13 AΒ The effects of the calcium channel blocker flunarizine on spinal monosynaptic reflexes were investigated in spinal cats. Flunarizine was administered locally into the spinal cord (10, 50, 100 .mu.M) and i.p. (5, 10, 20 mg/kg). Adult cats (n = 10) weighing 1.5-3kg were anesthetized with ketamine (50 mg/kg, i.m.), artificially ventilated and spinalized at the C1 level. A laminectomy was performed in the lumbosacral region. The ventral and dorsal roots of segment L5 were isolated and a pouch of skin formed at the site of the dissection to allow the exposed tissues to be covered with liq. paraffin. The temp. was kept at 38.5.degree.C with a heating pad. A polyethylene cannula was introduced into the left carotid artery to monitor blood pressure, which was kept above 100 mmHg. The dorsal root of segment L5 was placed on a silver-silver chloride wire electrode for stimulation through an isolation unit. The reflex potentials were recorded from the ipsilateral L5 ventral root, mounted on a silver-silver chloride wire electrode. The systemic (5, 10, 20 mg/kg) and local (50 and 100 .mu.M) dosages of flunarizine significantly decreased the amplitude of the reflex response (P < 0.05). Moreover, the latency of the monosynaptic reflexes increased after administration of the drug (P < 0.05). Voltage-dependent calcium channels in the spinal cord may play an important role in regulating the reflex response. ST flunarizine calcium channel spinal reflex antiepileptic ΙT Calcium channel RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (L-type; effects of calcium channel blocker

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL

flunarizine on spinal reflexes in cats)

(T-type; effects of calcium channel blocker

(Biological study); PROC (Process)

IT

Calcium channel

```
flunarizine on spinal reflexes in cats)
ΙT
     Ion channel blockers
        (calcium; effects of calcium channel
        blocker flunarizine on spinal reflexes in cats)
ΤТ
     Sodium channel
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (effects of calcium channel blocker
        flunarizine on spinal reflexes in cats)
TΤ
     Anticonvulsants
        (effects of calcium channel blocker
        flunarizine on spinal reflexes in cats in relation to epilepsy)
ΙT
     Reflex
        (spinal; effects of calcium channel blocker
        flunarizine on spinal reflexes in cats)
     52468-60-7, Flunarizine
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (effects of calcium channel blocker
        flunarizine on spinal reflexes in cats)
              THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE.
(1) Aicardi, G; Exp Brain Res 1990, V81, P288 CAPLUS
(2) Binnie, C; Ann NY Acad Sci 1988, V522, P710
(3) Carbone, E; J Physiol 1987, V386, P547 CAPLUS (4) Carbone, E; J Physiol 1987, V386, P571 CAPLUS
(5) Carbone, E; Nature 1984, V310, P501 CAPLUS
(6) de Sarro, G; Br J Pharmac 1988, V93, P247 CAPLUS
(7) Ferlinz, J; Ann Intern Med 1986, V105, P714 MEDLINE
(8) Fujii, S; J Pharmacol Exp Ther 1997, V280(3), P1187
(9) Gan'Shina, T; Eksp Klin Farmakol 1996, V59(5), P12 CAPLUS
(10) Genc, O; First European Congress of Pharmacology Pharmacological Research
    Abst Book 1995, P226
(11) Gould, R; Mol Pharmacol 1984, V25, P235 CAPLUS
(12) Katz, A; Am J Cardiol 1985, V55, P2B CAPLUS
(13) Kawasaki, K; Jap J Pharmac 1978, V28, P165 CAPLUS
(14) Leysen, J; The Pharmacological Basis of Migraine Therapy 1984, P255 CAPLUS
(15) McLean, M; Pol J Pharmacol Pharm 1987, V39, P513 MEDLINE
(16) Mikkelsen, E; Ugeskr Laeger 1995, V157(26), P3750 MEDLINE
(17) Moraidis; Neurosci Lett 1991, V129, P51 CAPLUS
(18) Nagano, N; Gen Pharmac 1988, V19, P789 CAPLUS
(19) Nowycky, M; Nature 1985, V31, P339
(20) Popoli, P; Arch Int Pharmacodyn Ther 1988, V292, P58 CAPLUS
(21) Straub, H; Epilepsy and inhibition 1992, P255
(22) Straub, H; Pflugers Arch 1991, Suppl1, PR14
(23) Tytgat, J; Naunyn Schmiedebergs Arch Pharmacol 1988, V337, P690 CAPLUS
(24) Weiner, D; Med Clin N Am 1988, V72, P83 CAPLUS
L5
     ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS
     1992:143854 CAPLUS
AN
     116:143854
DN
     Treatment of cutaneous hypersensitivity with topical calcium
ΤI
     channel blockers
     Sharpe, Richard J.; Arndt, Kenneth A.; Galli, Stephen J.
ΤN
     Beth Israel Hospital Assoc., USA
PA
SO
     S. African, 23 pp.
     CODEN: SFXXAB
DT
     Patent
     English
LΑ
     ICM A61K
IC
     ICS CO7D
CC
     1-7 (Pharmacology)
```

```
Section cross-reference(s): 63
FAN.CNT 1
    PATENT NO.
                    KIND DATE
                                         APPLICATION NO. DATE
     -----
                                         ______
                    A 19910925
    ZA 9006583
                                         ZA 1990-6583
                                                          19900720
PΙ
PRAI US 1989-396846
                          19890821
    Ca channel blockers are used topically for inhibition of cutaneous,
AB
    ocular, or mucosal hypersensitivity reactions, inflammation,
    hyperproliferation, or scarring. Mice were sensitized to 3% oxazolone (I)
    by applying I to the abdomen and hind footpad. On the day of treatment,
     each side of both ears were challenged with I. Nifedipine (II) was
     applied to each side of a given ear 1 h after challenge. II reduced the
     I-induced inflammation significantly after 24 h as compared to control.
ST
     calcium channel blocker skin
    hypersensitivity; inflammation inhibitor calcium channel
    blocker; hyperproliferation inhibitor calcium
     channel blocker; scarring inhibitor calcium
     channel blocker; nifedipine inflammation inhibitor
     Inflammation inhibitors
IT
        (calcium channel blockers)
ΙT
    Leukocyte
        (cutaneous accumulation of, inhibition of, in skin
       hypersensitivity reactions)
ΙT
     Fungicides and Fungistats
     Steroids, biological studies
     RL: BIOL (Biological study)
        (cutaneous hypersensitivity inhibition with topical calcium
        channel blockers and)
ΙT
    Lupus erythematosus
        (cutaneous, treatment of, with topical calcium
        channel blockers)
ΙT
     Burn
     Psoriasis
       Skin, disease
        (treatment of, with topical calcium channel
       blockers)
IΤ
    Dermatitis
        (allergic, contact, treatment of, with topical calcium
        channel blockers)
    Ulcer inhibitors
IΤ
        (aphthous, topical calcium channel blockers
TΤ
    Ion channel blockers
        (calcium, cutaneous hypersensitivity inhibition with topical)
IT
    Dermatitis
        (contact, treatment of, with topical calcium channel
       blockers)
ΙT
    Mucous membrane
        (disease, treatment of, with topical calcium channel
       blockers)
TΤ
    Connective tissue
        (disease, scleroderma, treatment of, with topical calcium
        channel blockers)
IT
    Vagina
        (disease, vaginitis, treatment of, with topical calcium
        channel blockers)
ΙT
     Skin, disease
        (hyperproliferation, treatment of, with topical calcium
        channel blockers)
     Skin, disease
TΤ
        (ichthyosis, treatment of, with topical calcium
        channel blockers)
TΤ
    Eye, disease
```

```
(inflammation, treatment of, with topical calcium
       channel blockers)
ΙT
    Skin, disease
        (lichen planus, treatment of, with topical calcium
        channel blockers)
ΙT
     Skin, neoplasm
        (mycosis fungoides, treatment of, with topical calcium
        channel blockers)
ΙT
     Skin, disease
        (pyoderma gangrenosum, treatment of, with topical calcium
        channel blockers)
IT
     Intestine, disease
        (rectum, inflammation, treatment of, with topical calcium
        channel blockers)
    Neurotransmitter antagonists
ΙT
        (serotoninergic, cutaneous hypersensitivity inhibition with topical
        calcium channel blockers and)
ΙT
    Acne
        (vulgaris, treatment of, with topical calcium channel
       blockers)
                          298-57-7
                                    6621-47-2, Perhexiline
                                                              21829-25-4,
ΙT
     52-53-9, Verapamil
                  22609-73-0, Niludipine
                                           39562-70-4, Nitrendipine
     Nifedipine
     42399-41-7 52468-60-7, Flunarizine
                                          55985-32-5, Nicardipine
                                        66085-59-4, Nimodipine
                                                                  72509-76-3,
     63675-72-9
                  64706-54-3, Bepridil
                  75530-68-6, Nilvadipine
                                            75695-93-1, Isradipine
     Felodipine
     119615-65-5, McN-6186
     RL: BIOL (Biological study)
        (cutaneous hypersensitivity inhibition with topical)
     50-55-5, Reserpine
                         129-03-3, Cyproheptadine
                                                    288-32-4D, Imidazole,
TΤ
                          749-02-0, Spiperone 1166-34-3, Cinanserine
               361-37-5
                                                  19794-93-5, Trazodone
     1400-61-9, Nystatin 1893-33-0, Pipamperone
                                24219-97-4, Mianserin
                                                        27220-47-9
     23593-75-1, Clotrimazole
     41621-49-2, Ciclopirox olamine 60634-51-7, LY 53857
                                                            61318-90-9
     64211-45-6, Oxiconazole 65277-42-1 65472-88-0, Naftifine
                84625-61-6, Itraconazole 85273-96-7 87051-43-2,
     Ketanserin
                  106266-06-2, Risperidone
                                             108674-87-9, LY 281067
     Ritanserin
     RL: BIOL (Biological study)
        (cutaneous hypersensitivity inhibition with topical calcium
        channel blockers and)
=> d 1-2 all, hitstr
    ANSWER 1 OF 2 CAPLUS COPYRIGHT 2002 ACS
L5
     1999:339238 CAPLUS
ΑN
DN
     131:139372
     Effects of the calcium channel blocker
ΤI
     flunarizine on spinal reflexes in cats
     Genc, O.; Demir, S.; Tasci, N.; Kaptanoglu, B.; Marangoz, C.
ΑU
     Department of Physiology, Faculty of Medicine, Pamukkale University,
CS
     Denizli, 20070, Turk.
     Medical Science Research (1999), 27(4), 247-250
SO
     CODEN: MSCREJ; ISSN: 0269-8951
PB
     Lippincott Williams & Wilkins
DT
     Journal
     English
LA
CC
     1-11 (Pharmacology)
     Section cross-reference(s): 13
AB
     The effects of the calcium channel blocker
     flunarizine on spinal monosynaptic reflexes were investigated in spinal
     cats. Flunarizine was administered locally into the spinal cord (10, 50,
     100 .mu.M) and i.p. (5, 10, 20 \text{ mg/kg}). Adult cats (n = 10) weighing 1.5-3
     kg were anesthetized with ketamine (50 mg/kg, i.m.), artificially
```

ventilated and spinalized at the C1 level. A laminectomy was performed in the lumbosacral region. The ventral and dorsal roots of segment L5 were isolated and a pouch of skin formed at the site of the dissection to allow the exposed tissues to be covered with liq. paraffin. The temp. was kept at 38.5.degree.C with a heating pad. A polyethylene cannula was introduced into the left carotid artery to monitor blood pressure, which was kept above 100 mmHg. The dorsal root of segment L5 was placed on a silver-silver chloride wire electrode for stimulation through an isolation unit. The reflex potentials were recorded from the ipsilateral L5 ventral root, mounted on a silver-silver chloride wire electrode. The systemic (5, 10, 20 mg/kg) and local (50 and 100 .mu.M) dosages of flunarizine significantly decreased the amplitude of the reflex response (P < 0.05). Moreover, the latency of the monosynaptic reflexes increased after administration of the drug (P < 0.05). Voltage-dependent calcium channels in the spinal cord may play an important role in regulating the reflex response. flunarizine calcium channel spinal reflex antiepileptic Calcium channel RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (L-type; effects of calcium channel blocker flunarizine on spinal reflexes in cats) Calcium channel RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (T-type; effects of calcium channel blocker flunarizine on spinal reflexes in cats) Ion channel blockers (calcium; effects of calcium channel blocker flunarizine on spinal reflexes in cats) Sodium channel RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (effects of calcium channel blocker flunarizine on spinal reflexes in cats) Anticonvulsants (effects of calcium channel blocker flunarizine on spinal reflexes in cats in relation to epilepsy) Reflex (spinal; effects of calcium channel blocker flunarizine on spinal reflexes in cats) **52468-60-7**, Flunarizine RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (effects of calcium channel blocker flunarizine on spinal reflexes in cats) THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT (1) Aicardi, G; Exp Brain Res 1990, V81, P288 CAPLUS (2) Binnie, C; Ann NY Acad Sci 1988, V522, P710 (3) Carbone, E; J Physiol 1987, V386, P547 CAPLUS (4) Carbone, E; J Physiol 1987, V386, P571 CAPLUS (5) Carbone, E; Nature 1984, V310, P501 CAPLUS (6) de Sarro, G; Br J Pharmac 1988, V93, P247 CAPLUS (7) Ferlinz, J; Ann Intern Med 1986, V105, P714 MEDLINE (8) Fujii, S; J Pharmacol Exp Ther 1997, V280(3), P1187 (9) Gan'Shina, T; Eksp Klin Farmakol 1996, V59(5), P12 CAPLUS (10) Genc, O; First European Congress of Pharmacology Pharmacological Research Abst Book 1995, P226 (11) Gould, R; Mol Pharmacol 1984, V25, P235 CAPLUS

ST

ΙT

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RE

(12) Katz, A; Am J Cardiol 1985, V55, P2B CAPLUS

(13) Kawasaki, K; Jap J Pharmac 1978, V28, P165 CAPLUS

```
(14) Leysen, J; The Pharmacological Basis of Migraine Therapy 1984, P255 CAPLUS
```

(15) McLean, M; Pol J Pharmacol Pharm 1987, V39, P513 MEDLINE

(16) Mikkelsen, E; Ugeskr Laeger 1995, V157(26), P3750 MEDLINE

(17) Moraidis; Neurosci Lett 1991, V129, P51 CAPLUS

(18) Nagano, N; Gen Pharmac 1988, V19, P789 CAPLUS

(19) Nowycky, M; Nature 1985, V31, P339

(20) Popoli, P; Arch Int Pharmacodyn Ther 1988, V292, P58 CAPLUS

(21) Straub, H; Epilepsy and inhibition 1992, P255

(22) Straub, H; Pflugers Arch 1991, Suppl1, PR14

(23) Tytgat, J; Naunyn Schmiedebergs Arch Pharmacol 1988, V337, P690 CAPLUS

(24) Weiner, D; Med Clin N Am 1988, V72, P83 CAPLUS

52468-60-7, Flunarizine ΙT

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(effects of calcium channel blocker

flunarizine on spinal reflexes in cats)

52468-60-7 CAPLUS
Piperazine, 1-[bis(4-fluorophenyl)methyl]-4-[(2E)-3-phenyl-2-propenyl]-CN (CA INDEX NAME)

Double bond geometry as shown.

```
ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS
L5
```

ΑN 1992:143854 CAPLUS

116:143854 DN

Treatment of cutaneous hypersensitivity with topical calcium TТ channel blockers

Sharpe, Richard J.; Arndt, Kenneth A.; Galli, Stephen J. IN

Beth Israel Hospital Assoc., USA PA

SO S. African, 23 pp.

CODEN: SFXXAB

DT Patent

English LA

IC ICM A61K

ICS CO7D

CC 1-7 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	ZA 9006583	A	19910925	ZA 1990-6583	19900720

PRAI US 1989-396846 19890821

Ca channel blockers are used topically for inhibition of cutaneous, ocular, or mucosal hypersensitivity reactions, inflammation, hyperproliferation, or scarring. Mice were sensitized to 3% oxazolone (I) by applying I to the abdomen and hind footpad. On the day of treatment,

```
each side of both ears were challenged with I. Nifedipine (II) was
     applied to each side of a given ear 1 h after challenge. II reduced the
     I-induced inflammation significantly after 24 h as compared to control.
ST
     calcium channel blocker skin
     hypersensitivity; inflammation inhibitor calcium channel
     blocker; hyperproliferation inhibitor calcium
     channel blocker; scarring inhibitor calcium
     channel blocker; nifedipine inflammation inhibitor
     Inflammation inhibitors
ΤT
        (calcium channel blockers)
ΙT
     Leukocyte
        (cutaneous accumulation of, inhibition of, in skin
        hypersensitivity reactions)
     Fungicides and Fungistats
IΤ
     Steroids, biological studies
     RL: BIOL (Biological study)
        (cutaneous hypersensitivity inhibition with topical calcium
        channel blockers and)
ΙT
     Lupus erythematosus
        (cutaneous, treatment of, with topical calcium
        channel blockers)
IT
     Burn
     Psoriasis
       Skin, disease
        (treatment of, with topical calcium channel
        blockers)
IΤ
     Dermatitis
        (allergic, contact, treatment of, with topical calcium
        channel blockers)
     Ulcer inhibitors
IT
        (aphthous, topical calcium channel blockers
ΙT
     Ion channel blockers
        (calcium, cutaneous hypersensitivity inhibition with topical)
IT
     Dermatitis
        (contact, treatment of, with topical calcium channel
        blockers)
ΙT
     Mucous membrane
        (disease, treatment of, with topical calcium channel
        blockers)
ΙT
     Connective tissue
        (disease, scleroderma, treatment of, with topical calcium
        channel blockers)
ΙT
     Vagina
        (disease, vaginitis, treatment of, with topical calcium
        channel blockers)
     Skin, disease
IΤ
        (hyperproliferation, treatment of, with topical calcium
        channel blockers)
     Skin, disease
IT
        (ichthyosis, treatment of, with topical calcium
        channel blockers)
TΤ
     Eye, disease
        (inflammation, treatment of, with topical calcium
        channel blockers)
     Skin, disease
IT
        (lichen planus, treatment of, with topical calcium
        channel blockers)
IT
     Skin, neoplasm
        (mycosis fungoides, treatment of, with topical calcium
        channel blockers)
ΙT
     Skin, disease
        (pyoderma gangrenosum, treatment of, with topical calcium
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channel blockers)
     Intestine, disease
ΙT
        (rectum, inflammation, treatment of, with topical calcium
        channel blockers)
ΙT
     Neurotransmitter antagonists
        (serotoninergic, cutaneous hypersensitivity inhibition with topical
        calcium channel blockers and)
ΙT
     Acne
        (vulgaris, treatment of, with topical calcium channel
        blockers)
                           298-57-7
                                       6621-47-2, Perhexiline
                                                                  21829-25-4,
TΤ
     52-53-9, Verapamil
     Nifedipine 22609-73-0, Niludipine 39562-70-4, Nitrendipine
     42399-41-7 52468-60-7, Flunarizine 55985-32-5, Nicardipine
                                                                     72509-76-3,
                   64706-54-3, Bepridil 66085-59-4, Nimodipine
     63675-72-9
                   75530-68-6, Nilvadipine 75695-93-1, Isradipine
     Felodipine
     119615-65-5, McN-6186
     RL: BIOL (Biological study)
        (cutaneous hypersensitivity inhibition with topical)
                         129-03-3, Cyproheptadine 288-32-4D, Imidazole, 749-02-0, Spiperone 1166-34-3, Cinanserine
ΙT
     50-55-5, Reserpine
     derivs. 361-37-5 749-02-0, Spiperone 1166-34-3, Cinanserine 1400-61-9, Nystatin 1893-33-0, Pipamperone 19794-93-5, Trazodone
                                24219-97-4, Mianserin 27220-47-9
     23593-75-1, Clotrimazole
     41621-49-2, Ciclopirox olamine
                                                               61318-90-9
                                        60634-51-7, LY 53857
     64211-45-6, Oxiconazole 65277-42-1 65472-88-0, Naftifine 74050-98-9,
     Ketanserin 84625-61-6, Itraconazole
                                               85273-96-7 87051-43-2,
                                               108674-87-9, LY 281067
                  106266-06-2, Risperidone
     Ritanserin
     RL: BIOL (Biological study)
        (cutaneous hypersensitivity inhibition with topical calcium
        channel blockers and)
     52468-60-7, Flunarizine
ΙT
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(cutaneous hypersensitivity inhibition with topical)

52468-60-7 CAPLUS
Piperazine, 1-[bis(4-fluorophenyl)methyl]-4-[(2E)-3-phenyl-2-propenyl]-

Double bond geometry as shown.

(9CI) (CA INDEX NAME)

RN CN

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RL: BIOL (Biological study)

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L10 ANSWER 1 OF 3 USPATFULL
IN
       Breton, Lionel, Versailles, FRANCE
       Nonotte, Isabelle, Paris, FRANCE
TI
       Treating skin wrinkles/fine lines with calcium channel
       inhibitors
                               20020516
PΙ
       US 2002058682
                          A 1
L10 ANSWER 2 OF 3 USPATFULL
ΙN
       Nonotte, Isabelle, Paris, FRANCE
       Breton, Lionel, Versailles, FRANCE
ΤI
       Manganese compositions for reducing/preventing skin wrinkles and fine
       lines
       US 2002028254
                               20020307
PΙ
                          A1
L10 ANSWER 3 OF 3 USPATFULL
       Breton, Lionel, Versailles, FRANCE
ΙN
       Nonotte, Isabelle, Paris, FRANCE
ΤI
       Treating skin wrinkles/fine lines with calcium channel
       inhibitors
                         B1 20020205
PΙ
       US 6344461
=> d his
     (FILE 'HOME' ENTERED AT 09:11:56 ON 31 MAY 2002)
     FILE 'CAPLUS' ENTERED AT 09:12:06 ON 31 MAY 2002
L1
             32 S VARIOUS CALCIUM CHANNEL
L2
          35687 S CALCIUM (5A) (INHIBIT? OR BLOCK? OR ANTAGONI?)
           6422 S (SMOOTH? OR RELAX? OR LOOSEN? OR SLACKEN?) (10A) (SKIN OR TIS
L3
             33 S L2 (2S) L3
L4
L5
             28 S L2 (1S) L3
L6
             22 S L5/AB
L7
             22 FOCUS L6 1-
     FILE 'USPATFULL' ENTERED AT 10:08:31 ON 31 MAY 2002
            105 S L5
L8
L9
             32 S (SMOOTH? OR RELAX? OR LOOSEN? OR SLACKEN?) (10A) (SKIN TISSUE
L10
              3 S L2 AND L9
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